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## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Inventors: **Chen W. Liaw et al.**Serial No.: **Not yet assigned**Group Art Unit: **1646**Filed: **Concurrently herewith**Examiner: **Prema Maria Mertz**Title: **NUCLEIC ACID ENCODING HUMAN G PROTEIN-COUPLED RECEPTOR****EXPRESS MAIL INFORMATION**

EXPRESS MAIL LABEL NO: EL964555223US

DATE OF DEPOSIT: February 19, 2004

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**INFORMATION DISCLOSURE STATEMENT****S I R :**

Pursuant to 37 C.F.R. §§ 1.97 and 1.98 and to the duty of disclosure set forth in 37 C.F.R. § 1.56, the Examiner in charge of the above-identified application is requested to consider and make of record the references listed on the attached PTO/SB/08a, PTO/SB/08b, PTO-1449 and PTO-892 forms submitted herewith.

Although the information submitted herewith may be "material" to the Examiner's consideration of the subject application, this submission is not intended to constitute an admission that such information is "prior art" as to the claimed invention.

Copies of the references cited on the attached PTO/SB/08a, PTO/SB/08b, PTO-1449 and PTO-892 forms can be found in the parent case, U.S. Serial No. 09/875,076, filed June 6, 2001.

In accordance with 37 C.F.R. § 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made.

No first Official Action has yet been received and it is presumed that none has yet been mailed. No fee or certification is required. 37 C.F.R. § 1.97(b).

Respectfully submitted,

A handwritten signature in dark ink, appearing to read "Michael A. Patané", is written over a horizontal line.

Michael A. Patané

Regis. No. 42,982

Enclosures:

24 sheets attached

Dated: 02/19/2004

Cozen O'Connor  
1900 Market Street  
Philadelphia, PA 19103  
Tel: 215.665.2000  
Fax: 215.665.2013

Substitute for form 1449A/PTO

## INFORMATION DISCLOSURE STATEMENT BY APPLICANT

*(use as many sheets as necessary)*

Sheet	1	of	2
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**Complete if Known**

Application Number	09/875,076
Filing Date	June 6, 2001
First Named Inventor	C. Liaw et al.
Art Unit	1646
Examiner Name	Prema Maria Mertz
Attorney Docket Number	AREN-11.US9.DIV (0239)

## U.S. PATENT DOCUMENTS

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This collection of information is required by 37 CFR 1.97 and 1.98. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 2 hours to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

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		Application Number	09/875,076
		Filing Date	June 6, 2001
		First Named Inventor	C. Liaw et al.
		Group Art Unit	1646
		Examiner Name	Prema Maria Mertz
Sheet 2 of 2	Attorney Docket Number	AREN-11.US9.DIV (0239)	

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		Hirsch et al., "Neuronal Vulnerability in Parkinson's Disease", <i>J Neural Transm Suppl.</i> 1997;50:79-88	
		Hulley et al., "Inhibitors of Type IV Phosphodiesterases Reduce the Toxicity of MPTP in Substantia Nigra Neurons <i>In Vivo</i> ", <i>European Journal of Neuroscience</i> , Vol. 7, pp. 2431-2440, 1995	
		Berridge, Michael J., "Inositol trisphosphate and calcium signalling", <i>Nature</i> , Vol. 361, 28 January 1993	
		Wilson et al., "Orphan G-Protein-coupled receptors: the next generation of drug targets?", <i>British Journal of Pharmacology</i> (1998) 125, 1387-1392	
		Hegyi et al., "The Relationship between Protein Structure and Function: a Comprehensive Survey with Application to the Yeast Genome", <i>J. Mol. Biol.</i> (1999) 288, 147-164	
		Kasuya et al., "Three-dimensional Structure Analysis of PROSITE Patterns", <i>J. Mol. Biol.</i> (1999) 288, 1673-1691	

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\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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**INFORMATION DISCLOSURE  
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Sheet 2 of 2

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Application Number	09/875,076
Filing Date	June 6, 2001
First Named Inventor	Chen Liaw
Group Art Unit	1646
Examiner Name	P. Mertz
Attorney Docket Number	11.US9.DIV (AREN-0239)

**OTHER PRIOR ART – NON PATENT LITERATURE DOCUMENTS**

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	BA	Standaert and Young, Treatment of Central Nervous System Degenerative Disorders, in Goodman & Gilman's The Pharmaceutical Basis of Therapeutics, Ninth Ed. (Chapter 22) pp. 503-519, 1996	
	BB	Matsumoto, et al., "An Evolutionarily Conserved G-Protein Coupled receptor Family, SREB, Expressed in the Central Nervous System," Biochem. Biophys. Res. Comm. (2000) 272:576-582	
	BC	Skolnick, et al., "Structural genomics and its importance for gene function analysis," Nature Biotechnol. (2000) 18:283-287	

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<b>Form PTO-1449 Modified</b>  List of Patent and Publications Cited by Applicant (Use several sheets if necessary)  U.S. Department of Commerce Patent and Trademark Office		Docket No. <b>AREN-0239</b>	Serial No. <b>09/875,076</b>
		Applicant <b>Ruoping Chen, et al.</b>	
		Filing Date <b>June 6, 2001</b>	Group <b>Not Yet Assigned</b>
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	<b>AA</b>	Alla, S.A. et al., "Extracellular domains of the bradykinin B2 receptor involved in ligand binding and agonist sensing defined by anti-peptide antibodies," <i>J. Biol. Chem.</i> , 1996, 271, 1748-1755	
	<b>AB</b>	Advenier, C. et al., "Effects on the isolated human bronchus of SR 48968, a potent and selective nonpeptide antagonist of the neurokinin A (NK <sub>2</sub> ) receptors," <i>Am. Rev. Respir. Dis.</i> , 1992, 146(5, Pt. 1), 1177-1181	
	<b>AC</b>	Alexander, W.S. et al., "Point mutations within the dimer interfact homology domain of c-Mpl induce constitutive receptor activity and tumorigenicity," <i>EMBO J.</i> , 1995, 14(22), 5569-5578	
	<b>AD</b>	Arvanitakis, L. et al., "Human herpesvirus KSHV encodes a constitutively active G-protein-coupled receptor linked to cell proliferation," <i>Nature</i> , 1997, 385, 347-349	
	<b>AE</b>	Barker, E.L. et al., "Constitutively active 5-hydroxytryptamine <sub>2C</sub> receptors reveal novel inverse agonist activity of receptor ligands," <i>J. Biol. Chem.</i> , 1994, 269(16), 11687-11690	
	<b>AF</b>	Baxter, G., "5-HT <sub>2</sub> receptors: a family re-united?" <i>Trends Pharmacol. Sci.</i> , 1995, 16, 105-110	
	<b>AG</b>	Besmer, P. et al., "A new acute transforming feline retrovirus and relationship of its oncogene <i>v-kit</i> with the protein kinase gene family," <i>Nature</i> , 1986, 320, 415	
	<b>AH</b>	Blin, N. et al., "Mapping of single amino acid residues required for selective activation of G <sub>q/11</sub> by the m3 muscarinic acetylcholine receptor," <i>J. Biol. Chem.</i> , 1995, 270, 17741-17748	
	<b>AI</b>	Bond, R.A. et al., "Inverse agonists and G-protein-coupled receptors," in <i>Receptor-Based Drug Design</i> , Leff, P. (ed.), New York, M. Dekker, 1998, 363-377	
	<b>AJ</b>	Boone, C. et al., "Mutations that alter the third cytoplasmic loop of the a-factor receptor lead to a constitutive and hypersensitive phenotype," <i>Proc. Natl. Acad. Sci. USA</i> , 1993, 90(21), 9921-9925	
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	<b>AK</b>	Burstein, E.S. et al., "Constitutive activation of chimeric m2/m5 muscarinic receptors and delineation of G-protein coupling selectivity domains," <i>Biochem. Pharmacol.</i> , 1996, 51(4), 539-544	
	<b>AL</b>	Burstein, E.S. et al., "Amino acid side chains that define muscarinic receptor/G-protein coupling. Studies of the third intracellular loop," <i>J. Biol. Chem.</i> , 1996, 271(6), 2882-2885	
	<b>AM</b>	Burstein, E.S. et al., "Constitutive activation of muscarinic receptors by the G-protein G <sub>q</sub> ," <i>FEBS Lett.</i> , 1995, 363(3), 261-263	
	<b>AN</b>	Bylund, D., "International union of pharmacology nomenclature of adrenoceptors," <i>Pharmacol. Rev.</i> , 1994, 46, 121-136	
	<b>AO</b>	Casey, C. et al., "Constitutively active mutant 5-HT <sub>2A</sub> serotonin receptors: inverse agonist activity of classical 5HT <sub>2A</sub> antagonists," <i>Soc. Neurosci.</i> , 1996, Abstract #699.10	
	<b>AP</b>	Cheatham, B. et al., "Substitution of the <i>erbB-2</i> oncoprotein transmembrane domain activates the insulin receptor and modulates the action of insulin-receptor substrate 1," <i>Proc. Natl. Acad. Sci. USA</i> , 1993, 90, 7336-7340	
	<b>AQ</b>	Chen, J. et al., "Tethered Ligand Library for Discovery of Peptide Agonists," <i>J. Biol. Chem.</i> , 1995, 270, 23398-23401	
	<b>AR</b>	Chen, T.S. et al., "Microbial hydroxylation and glucuronidation of the angiotensin II (AII) receptor antagonist MK 954," <i>J. Antibiot. (Tokyo)</i> , 1993, 46(1), 131-134	
	<b>AS</b>	Chen, W. et al., "A colorimetric assay for measuring activation of G <sub>s</sub> - and G <sub>q</sub> -coupled signaling pathways," <i>Anal. Biochem.</i> , 1995, 226(2), 349-354	
	<b>AT</b>	Chidiac, P. et al., "Inverse agonist activity of $\beta$ -adrenergic antagonists," <i>J. Pharm. Exp. Ther.</i> , 1994, 45, 490-499	
	<b>AU</b>	Clozel, M. et al., "In vivo pharmacology of Ro 46-2005, the first synthetic nonpeptide endothelin receptor antagonist: implications for endothelin physiology," <i>J. Cardiovas. Pharmacol.</i> , 1993, 22(Suppl. 8), S377-S379	
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<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	<b>AV</b>	Collesi, C. et al., "A splicing variant of the <i>RON</i> transcript induces constitutive tyrosine kinase activity and an invasive phenotype," <i>Mol. Cell. Biol.</i> , 1996, 16(2), 5518-5526	
	<b>AW</b>	Cooper, C.S. et al., "Molecular cloning of a new transforming gene from a chemically transformed human cell line," <i>Nature</i> , 1984, 311, 29-33	
	<b>AX</b>	De Dios, I. et al., "Effect of L-364,718 (CCK Receptor Antagonist) on Exocrine Pancreatic Secretion of Hydrocortison-Treated Rats," <i>Pancreas</i> , 1994, 9(2), 212-218	
	<b>AY</b>	Desbios-Mouthon, C. et al., "Deletion of Asn <sup>281</sup> in the $\alpha$ -subunit of the human insulin receptor causes constitutive activation of the receptor and insulin desensitization," <i>J. Clin. Endocrinol. Metab.</i> , 1996, 81(2), 719-727	
	<b>AZ</b>	Di Renzo, M.F. et al., "Expression of the Met/HGF receptor in normal and neoplastic human tissues," <i>Oncogene</i> , 1991, 6(11), 1997-2003	
	<b>BA</b>	Di Renzo, M.F. et al., "Overexpression of the c-MET/HGF receptor gene in human thyroid carcinomas," <i>Oncogene</i> , 1992, 7, 2549-2553	
	<b>BB</b>	Duprez, L. et al., "Germline mutations of the thyrotropin receptor gene cause non-autoimmune autosomal dominant hyperthyroidism," <i>Nature Genetics</i> , 1994, 7, 396-401	
	<b>BC</b>	Eggericksx, D. et al., "Molecular Cloning of an Orphan G-Protein-Coupled Receptor that Constitutively Activates Adenylate Cyclase," <i>Biochem. J.</i> , 1995, 309, 837-843	
	<b>BD</b>	Evans, B.E. et al., "Orally Active, Nonpeptide Oxytocin Antagonists," <i>J. Med. Chem.</i> , 1992, 35, 3919-3927	
	<b>BE</b>	Fu, M. et al., "Functional autoimmune epitope on $\alpha_1$ -adrenergic receptors in patients with malignant hypertension," <i>Lancet</i> , 1994, 344, 1660-1663	
	<b>BF</b>	Furitsu, T. et al., "Identification of Mutations in the Coding Sequence of the Proto-oncogene <i>c-kit</i> in a Human Mast Cell Leukemia Cell Line Causing Ligand-independent Activation of <i>c-kit</i> Product," <i>J. Clin. Invest.</i> , 1993, 92, 1736-1744	
	<b>BG</b>	Gellai, M. et al., "Nonpeptide Endothelin Receptor Antagonists V: Prevention and Reversal of Acute Renal Failure in the Rat by SB 209670," <i>J. Pharm. Exp. Therap.</i> , 1995, 275(1), 200-206	
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<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	<b>BH</b>	Gitter, B. et al., "Pharmacological Characterization of LY303870: A Novel Potent and Selective Nonpeptide Substance P (Neurokinin-1) Receptor Antagonist," <i>J. Pharm. Exp. Therp.</i> , <b>1995</b> , 275(2), 737-744	
	<b>BI</b>	Gouilleux-Gruart, V. et al., "STAT-Related Transcription Factors are Constitutively Activated in Peripheral Blood Cells from Acute Leukemia Patients," <i>Blood</i> , <b>1996</b> , 87(5), 1692-1697	
	<b>BJ</b>	Hansson, J.H. et al., "Hypertension caused by a truncated epithelial sodium channel $\gamma$ subunit: genetic heterogeneity of Liddle syndrome," <i>Nat. Genet.</i> , <b>1995</b> , 11(1), 76-82	
	<b>BK</b>	Hasegawa, H. et al., "Two Isoforms of the Prostaglandin E Receptor EP3 Subtype Different in Agonist-independent Constitutive Activity," <i>J. Biol. Chem.</i> , <b>1996</b> , 271(4), 1857-1860	
	<b>BL</b>	Hendler, F. et al., "Human Squamous Cell Lung Cancers Express Increased Epidermal Growth Factor Receptors," <i>J. Clin. Invest.</i> , <b>1984</b> , 74, 647-651	
	<b>BM</b>	Herrick-Davis, K. et al., "Constitutively Active 5HT <sub>2C</sub> Serotonin Receptor Created by Site-Directed Mutagenesis," <i>Soc. Neurosci.</i> , Abstract No. 699.18	
	<b>BN</b>	Hieble, J., "International union of pharmacology. X. Recommendation for nomenclature of 1-adrenoceptors," <i>Pharm. Rev.</i> , <b>1995</b> , 47, 267-270	
	<b>BO</b>	Hill, S., "Distribution, Properties, and Functional Characteristics of Three Classes of Histamine Receptor," <i>Am. Soc. Pharm. Exp. Therap.</i> , <b>1990</b> , 42(1), 45-83	
	<b>BP</b>	Högger, P. et al., "Activating and Inactivating Mutations in - and C-terminal i3 Loop Junctions of Muscarinic Acetylcholine Hm1 Receptors," <i>J. Biol. Chem.</i> , <b>1995</b> , 270(13), 7405-7410	
	<b>BQ</b>	Ikeda, H. et al., "Expression and Functional Role of the Proto-oncogene <i>c-kit</i> in Acute Myeloblastic Leukemia Cells," <i>Blood</i> , <b>1991</b> , 78(11), 2962-2968	
	<b>BR</b>	Imura, R. et al., "Inhibition by HS-142-1, a novel nonpeptide atrial natriuretic peptide antagonist of microbial origin, of atrial natriuretic peptide-induced relaxation of isolated rabbit aorta through the blockade of guanylyl cyclase-linked receptors," <i>Mol. Pharm.</i> , <b>1992</b> , 42, 982-990	
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<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>			
	<b>BS</b>	Jakubik, J. et al., "Constitutive activity of the M <sub>1</sub> -M <sub>4</sub> subtypes of muscarinic receptors in transfected CHO cells and of muscarinic receptors in the heart cells revealed by negative antagonists," <i>FEBS Letts.</i> , 1995, 377, 275-279	
	<b>BT</b>	Kjelsberg, M.A. et al., "Constitutive activation of the $\alpha_{1B}$ -adrenergic receptor by all amino acid substitutions at a single site," <i>J. Biol. Chem.</i> , 1992, 267(3), 1430-1433	
	<b>BU</b>	Knapp, R. et al., "Molecular biology and pharmacology of cloned opioid receptors," <i>FASEB J.</i> , 1995, 9, 516-525	
	<b>BV</b>	Kosugi, S. et al., "Characterization of heterogeneous mutations causing constitutive activation of the luteinizing hormone receptor in familial male precocious puberty," <i>Human Mol. Genetics</i> , 1995, 4(2), 183-188	
	<b>BW</b>	Kosugi, S. et al., "Identification of Thyroid-Stimulating Antibody-Specific Interaction Sites in the N-Terminal Region of the Thyrotropin Receptor," <i>Mol. Endocrinology</i> , 1993, 7, 114-130	
	<b>BX</b>	Kraus, M. et al., "Demonstration of ligand-dependent signaling by the <i>erbB-3</i> tyrosine kinase and its constitutive activation in human breast tumor cells," <i>Proc. Natl. Acad. Sci. USA</i> , 1993, 90, 2900-2904	
	<b>BY</b>	Kudlacz, E. et al., "In Vitro and In Vivo Characterization of MDL 105,212A, a Nonpeptide NK-1/NK-2 Tachykinin Receptor Antagonist," <i>J. Pharm. Exp. Therap.</i> , 1996, 277(2), 840-851	
	<b>BZ</b>	Kuriu, A. et al., "Proliferation of Human Myeloid Leukemia Cell Line Associated with the Tyrosine-Phosphorylation and Activation of the Proto-oncogene <i>c-kit</i> Product," <i>Blood</i> , 1991, 78(11), 2834-2840	
	<b>CA</b>	Labbé-Jullié, C. et al., "Effect of the nonpeptide neurotensin antagonist, SR 48692, and two enantiomeric analogs, SR 48527 and SR 49711, on neurotension binding and contractile responses in guinea pig ileum and colon," <i>J. Pharm. Exp. Therap.</i> , 1994, 271(1), 267-276	
	<b>CB</b>	Latronico, A. et al., "A novel mutation of the luteinizing hormone receptor gene causing male gonadotropin-independent precocious puberty," <i>J. Clin. Endocrinol. Metab.</i> , 1995, 80(8), 2490-2494	
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	<b>CC</b>	Laue, L. et al., "Genetic heterogeneity of constitutively activating mutations of the human luteinizing hormone receptor in familial male-limited precocious puberty," <i>Proc. Natl. Acad. Sci USA</i> , 1995, 92, 1906-1910	
	<b>CD</b>	Løvlie, R. et al., "The Ca <sup>2+</sup> -sensing receptor gene (PCAR1) mutation T151M in isolated autosomal dominant hypoparathyroidism," <i>Hum. Genet.</i> , 1996, 98, 129-133	
	<b>CE</b>	Lefkowitz, R. et al., "Constitutive activity of receptors coupled to guanine nucleotide regulatory proteins," <i>Trends Pharmacol. Sci.</i> , 1993, 14, 300-307	
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	<b>CG</b>	Liu, C. et al., "Overexpression of c-met proto-oncogene but not epidermal growth factor receptor or c-erbB-2 in primary human colorectal carcinomas," <i>Oncogene</i> , 1992, 7, 181-185	
	<b>CH</b>	Liu, J. et al., "Molecular mechanisms involved in muscarinic acetylcholine receptor-mediated G protein activation studied by insertion mutagenesis," <i>J. Biol. Chem.</i> , 1996, 271(11), 6172-6178	
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				Applicant <b>Ruoping Chen, et al.</b>			
				Filing Date <b>June 6, 2001</b>		Group <b>Not Yet Assigned</b>	
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	<b>GJ</b>	5,532,157	07/02/96	Fink	435	240.2	
	<b>GK</b>	5,573,944	11/12/96	Kirschner et al.	435	252.3	
	<b>GL</b>	5,639,616	06/17/97	Liao et al.	435	7.1	
	<b>GM</b>	5,750,353	05/12/98	Kopin et al.	435	7.21	
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Examiner Initial		Document No.	Date	Country	Translation YES NO		
	<b>GN</b>	WO 97/11159	09/20/96	PCT	X		
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\* A copy of this reference is not being furnished with this Office action.  
(See Manual of Patent Examining Procedure, Section 707.05(a).)